

SEQUENCE LISTING

<110> Raghuram Kalluri

<120> ANTI-ANGIOGENIC PROTEINS AND FRAGMENTS
AND METHODS OF USE THEREOF

<130> 1440.1027-016

<150> PCT/US01/00565

<151> 2001-01-08

<150> US 09/543,371

<151> 2000-04-04

<150> US 09/335,224

<151> 1999-06-17

<150> US 60/126,175

<151> 1999-03-25

<150> US 60/089,689

<151> 1998-06-17

<150> US 09/479,118

<151> 2000-01-07

<150> US 09/625,191

<151> 2000-07-21

<160> 58

<170> FastSEQ for Windows Version 4.0

<210> 1

<211> 690

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> (1)...(687)

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gac	cca	cag	tgt	cct	tct	ggg	acc	aaa	att	ctt	tac	cac	ggg	tac	tct	96
Asp	Pro	Gln	Cys	Pro	Ser	Gly	Thr	Lys	Ile	Leu	Tyr	His	Gly	Tyr	Ser	
		20						25					30			

ttg	ctc	tac	gtg	caa	ggc	aat	gaa	cgg	gcc	cat	gga	cag	gac	ttg	ggc	144
Leu	Leu	Tyr	Val	Gln	Gly	Asn	Glu	Arg	Ala	His	Gly	Gln	Asp	Leu	Gly	
		35					40					45				

acg gcc ggc agc tgc ctg cgc aag ttc agc aca atg ccc ttc ctg ttc	192
Thr Ala Gly Ser Cys Leu Arg Lys Phe Ser Thr Met Pro Phe Leu Phe	
50 55 60	
tgc aat att aac aac gtg tgc aac ttt gca tca cga aat gac tac tcg	240
Cys Asn Ile Asn Asn Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser	
65 70 75 80	
tac tgg ctg tcc acc cct gag ccc atg ccc atg tca atg gca ccc atc	288
Tyr Trp Leu Ser Thr Pro Glu Pro Met Pro Met Ser Met Ala Pro Ile	
85 90 95	
acg ggg gaa aac ata aga cca ttt att agt agg tgt gct gtg tgt gag	336
Thr Gly Glu Asn Ile Arg Pro Phe Ile Ser Arg Cys Ala Val Cys Glu	
100 105 110	
gcg cct gcc atg gtg atg gcc gtg cac agc cag acc att cag atc cca	384
Ala Pro Ala Met Val Met Ala Val His Ser Gln Thr Ile Gln Ile Pro	
115 120 125	
ccg tgc ccc agc ggg tgg tcc tcg ctg tgg atc ggc tac tct ttt gtg	432
Pro Cys Pro Ser Gly Trp Ser Ser Leu Trp Ile Gly Tyr Ser Phe Val	
130 135 140	
atg cac acc agc gct ggt gca gaa ggc tct ggc caa gcc ctg gcg tcc	480
Met His Thr Ser Ala Gly Ala Glu Gly Ser Gly Gln Ala Leu Ala Ser	
145 150 155 160	
ccc ggc tcc tgc ctg gag gag ttt aga agt gcg cca ttc atc gag tgt	528
Pro Gly Ser Cys Leu Glu Glu Phe Arg Ser Ala Pro Phe Ile Glu Cys	
165 170 175	
cac ggc cgt ggg acc tgc aat tac tac gca aac gct tac agc ttt tgg	576
His Gly Arg Gly Thr Cys Asn Tyr Tyr Ala Asn Ala Tyr Ser Phe Trp	
180 185 190	
ctc gcc acc ata gag agg agc gag atg ttc aag aag cct acg ccg tcc	624
Leu Ala Thr Ile Glu Arg Ser Glu Met Phe Lys Lys Pro Thr Pro Ser	
195 200 205	
acc ttg aag gca ggg gag ctg cgc acg cac gtc agc cgc tgc caa gtc	672
Thr Leu Lys Ala Gly Glu Leu Arg Thr His Val Ser Arg Cys Gln Val	
210 215 220	
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Cys Met Arg Arg Thr	
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<210> 2
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 <212> PRT
 <213> Homo sapiens

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 35 40 45
 Thr Ala Gly Ser Cys Leu Arg Lys Phe Ser Thr Met Pro Phe Leu Phe
 50 55 60
 Cys Asn Ile Asn Asn Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser
 65 70 75 80
 Tyr Trp Leu Ser Thr Pro Glu Pro Met Pro Met Ser Met Ala Pro Ile
 85 90 95
 Thr Gly Glu Asn Ile Arg Pro Phe Ile Ser Arg Cys Ala Val Cys Glu
 100 105 110
 Ala Pro Ala Met Val Met Ala Val His Ser Gln Thr Ile Gln Ile Pro
 115 120 125
 Pro Cys Pro Ser Gly Trp Ser Ser Leu Trp Ile Gly Tyr Ser Phe Val
 130 135 140
 Met His Thr Ser Ala Gly Ala Glu Gly Ser Gly Gln Ala Leu Ala Ser
 145 150 155 160
 Pro Gly Ser Cys Leu Glu Glu Phe Arg Ser Ala Pro Phe Ile Glu Cys
 165 170 175
 His Gly Arg Gly Thr Cys Asn Tyr Tyr Ala Asn Ala Tyr Ser Phe Trp
 180 185 190
 Leu Ala Thr Ile Glu Arg Ser Glu Met Phe Lys Lys Pro Thr Pro Ser
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<223> pET22b(+) forward oligonucleotide primer for Arresten

<400> 3

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<211> 27

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<213> Artificial Sequence

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<223> pET22b(+) reverse oligonucleotide primer for Arresten

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<211> 684

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<213> Homo sapiens

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<221> CDS

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ccc atg tgc ccg gtg ggc atg aac aaa ctc tgg agt gga tac agc ctg 96
 Pro Met Cys Pro Val Gly Met Asn Lys Leu Trp Ser Gly Tyr Ser Leu
 20 25 30

ctg tac ttc gag ggc cag gag aag gcg cac aac cag gac ctg ggg ctg 144
 Leu Tyr Phe Glu Gly Gln Glu Lys Ala His Asn Gln Asp Leu Gly Leu
 35 40 45

gcg ggc tcc tgc ctg gcg cgg ttc agc acc atg ccc ttc ctg tac tgc 192
 Ala Gly Ser Cys Leu Ala Arg Phe Ser Thr Met Pro Phe Leu Tyr Cys
 50 55 60

aac cct ggt gat gtc tgc tac tat gcc agc cgg aac gac aag tcc tac 240
 Asn Pro Gly Asp Val Cys Tyr Tyr Ala Ser Arg Asn Asp Lys Ser Tyr
 65 70 75 80

tgg ctc tct acc act gcg ccg ctg ccc atg atg ccc gtg gcc gag gac 288
 Trp Leu Ser Thr Thr Ala Pro Leu Pro Met Met Pro Val Ala Glu Asp
 85 90 95

gag atc aag ccc tac atc agc cgc tgt tct gtg tgt gag gcc ccg gcc 336
 Glu Ile Lys Pro Tyr Ile Ser Arg Cys Ser Val Cys Glu Ala Pro Ala
 100 105 110

atc gcc atc gcg gtc cac agt cag gat gtc tcc atc cca cac tgc cca 384
 Ile Ala Ile Ala Val His Ser Gln Asp Val Ser Ile Pro His Cys Pro
 115 120 125

gct ggg tgg cgg agt ttg tgg atc gga tat tcc ttc ctc atg cac acg 432
 Ala Gly Trp Arg Ser Leu Trp Ile Gly Tyr Ser Phe Leu Met His Thr
 130 135 140

gcg gcg gga gac gaa ggc ggt ggc caa tca ctg gtg tca ccg ggc agc 480
 Ala Ala Gly Asp Glu Gly Gly Gln Ser Leu Val Ser Pro Gly Ser
 145 150 155 160

tgt cta gag gac ttc cgc gcc aca cca ttc atc gaa tgc aat gga ggc 528
 Cys Leu Glu Asp Phe Arg Ala Thr Pro Phe Ile Glu Cys Asn Gly Gly
 165 170 175

cgc ggc acc tgc cac tac tac gcc aac aag tac agc ttc tgg ctg acc 576
 Arg Gly Thr Cys His Tyr Tyr Ala Asn Lys Tyr Ser Phe Trp Leu Thr
 180 185 190

acc att ccc gag cag agc ttc cag ggc tgc ccc tcc gcc gac acg ctc 624
 Thr Ile Pro Glu Gln Ser Phe Gln Gly Ser Pro Ser Ala Asp Thr Leu
 195 200 205

aag gcc ggc ctc atc cgc aca cac atc agc cgc tgc cag gtg tgc atg 672
 Lys Ala Gly Leu Ile Arg Thr His Ile Ser Arg Cys Gln Val Cys Met
 210 215 220

aag aac ctg tga 684
 Lys Asn Leu

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<210> 6
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 <212> PRT
 <213> Homo sapiens

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 Pro Met Cys Pro Val Gly Met Asn Lys Leu Trp Ser Gly Tyr Ser Leu
 20 25 30
 Leu Tyr Phe Glu Gly Gln Glu Lys Ala His Asn Gln Asp Leu Gly Leu
 35 40 45
 Ala Gly Ser Cys Leu Ala Arg Phe Ser Thr Met Pro Phe Leu Tyr Cys
 50 55 60
 Asn Pro Gly Asp Val Cys Tyr Tyr Ala Ser Arg Asn Asp Lys Ser Tyr
 65 70 75 80
 Trp Leu Ser Thr Thr Ala Pro Leu Pro Met Met Pro Val Ala Glu Asp
 85 90 95
 Glu Ile Lys Pro Tyr Ile Ser Arg Cys Ser Val Cys Glu Ala Pro Ala
 100 105 110
 Ile Ala Ile Ala Val His Ser Gln Asp Val Ser Ile Pro His Cys Pro
 115 120 125
 Ala Gly Trp Arg Ser Leu Trp Ile Gly Tyr Ser Phe Leu Met His Thr
 130 135 140
 Ala Ala Gly Asp Glu Gly Gly Gly Gln Ser Leu Val Ser Pro Gly Ser
 145 150 155 160
 Cys Leu Glu Asp Phe Arg Ala Thr Pro Phe Ile Glu Cys Asn Gly Gly
 165 170 175
 Arg Gly Thr Cys His Tyr Tyr Ala Asn Lys Tyr Ser Phe Trp Leu Thr
 180 185 190
 Thr Ile Pro Glu Gln Ser Phe Gln Gly Ser Pro Ser Ala Asp Thr Leu
 195 200 205
 Lys Ala Gly Leu Ile Arg Thr His Ile Ser Arg Cys Gln Val Cys Met
 210 215 220
 Lys Asn Leu

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 <210> 7
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 <212> DNA
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<220>
 <223> pET22b(+) forward oligonucleotide primer for
 Canstatin

<400> 7
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<210> 8
 <211> 27
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> pET22b(+) reverse oligonucleotide primer for
 Canstatin

<400> 8
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<210> 9
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<212> DNA
<213> Homo sapiens

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Thr Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile
20 25 30

cct tca tgt cca gag ggg aca gtg cca ctc tac agt ggg ttt tct ttt 144
Pro Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe
35 40 45

ctt ttt gta caa gga aat caa cga gcc cac gga caa gac ctt gga act 192
Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr
50 55 60

ctt ggc agc tgc ctg cag cga ttt acc aca atg cca ttc tta ttc tgc 240
Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys
65 70 75 80

aat gtc aat gat gta tgt aat ttt gca tct cga aat gat tat tca tac 288
Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr
85 90 95

tgg ctg tca aca cca gct ctg atg cca atg aac atg gct ccc att act 336
Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr
100 105 110

ggc aga gcc ctt gag cct tat ata agc aga tgc act gtt tgt gaa ggt 384
Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly
115 120 125

cct gcg atc gcc ata gcc gtt cac agc caa acc act gac att cct cca 432
Pro Ala Ile Ala Ile Ala Val His Ser Gln Thr Thr Asp Ile Pro Pro
130 135 140

tgt cct cac ggc tgg att tct ctc tgg aaa gga ttt tca ttc atc atg 480
Cys Pro His Gly Trp Ile Ser Leu Trp Lys Gly Phe Ser Phe Ile Met
145 150 155 160

ttc aca agt gca ggt tct gag ggc acc ggg caa gca ctg gcc tcc cct 528
Phe Thr Ser Ala Gly Ser Glu Gly Thr Gly Gln Ala Leu Ala Ser Pro
165 170 175

ggc tcc tgc ctg gaa gaa ttc cga gcc agc cca ttt cta gaa tgt cat 576
Gly Ser Cys Leu Glu Glu Phe Arg Ala Ser Pro Phe Leu Glu Cys His

cccaagcttc aggttcttca tgcacac
cca ggt ttg aaa gga aaa cgt gga gac agt gga tca cct gca acc tgg
Pro Gly Leu Lys Gly Lys Arg Gly Asp Ser Gly Ser Pro Ala Thr Trp
1 5 10 15
aca acg aga ggc ttt gtc ttc acc cga cac agt caa acc aca gca att
Thr Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile
20 25 30
cct tca tgt cca gag ggg aca gtg cca ctc tac agt ggg ttt tct ttt
Pro Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe
35 40 45
ctt ttt gta caa gga aat caa cga gcc cac gga caa gac ctt gga act
Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr
50 55 60
ctt ggc agc tgc ctg cag cga ttt acc aca atg cca ttc tta ttc tgc
Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys
65 70 75 80
aat gtc aat gat gta tgt aat ttt gca tct cga aat gat tat tca tac
Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr
85 90 95
tgg ctg tca aca cca gct ctg atg cca atg aac atg gct ccc att act
Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr
100 105 110
ggc aga gcc ctt gag cct tat ata agc aga tgc act gtt tgt gaa ggt
Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly
115 120 125
cct gcg atc gcc ata gcc gtt cac agc caa acc act gac att cct cca
Pro Ala Ile Ala Ile Ala Val His Ser Gln Thr Thr Asp Ile Pro Pro
130 135 140
tgt cct cac ggc tgg att tct ctc tgg aaa gga ttt tca ttc atc atg
Cys Pro His Gly Trp Ile Ser Leu Trp Lys Gly Phe Ser Phe Ile Met
145 150 155 160
ttc aca agt gca ggt tct gag ggc acc ggg caa gca ctg gcc tcc cct
Phe Thr Ser Ala Gly Ser Glu Gly Thr Gly Gln Ala Leu Ala Ser Pro
165 170 175
ggc tcc tgc ctg gaa gaa ttc cga gcc agc cca ttt cta gaa tgt cat
Gly Ser Cys Leu Glu Glu Phe Arg Ala Ser Pro Phe Leu Glu Cys His

<210> 10
<211> 245
<212> PRT
<213> Homo sapiens

<400> 10
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Thr Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile
20 25 30
Pro Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe
35 40 45
Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr
50 55 60
Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys
65 70 75 80
Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr
85 90 95
Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr
100 105 110
Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly
115 120 125
Pro Ala Ile Ala Ile Ala Val His Ser Gln Thr Thr Asp Ile Pro Pro
130 135 140
Cys Pro His Gly Trp Ile Ser Leu Trp Lys Gly Phe Ser Phe Ile Met
145 150 155 160
Phe Thr Ser Ala Gly Ser Glu Gly Thr Gly Gln Ala Leu Ala Ser Pro
165 170 175
Gly Ser Cys Leu Glu Glu Phe Arg Ala Ser Pro Phe Leu Glu Cys His
180 185 190
Gly Arg Gly Thr Cys Asn Tyr Tyr Ser Asn Ser Tyr Ser Phe Trp Leu
195 200 205
Ala Ser Leu Asn Pro Glu Arg Met Phe Arg Lys Pro Ile Pro Ser Thr
210 215 220
Val Lys Ala Gly Glu Leu Glu Lys Ile Ile Ser Arg Cys Gln Val Cys
225 230 235 240
Met Lys Lys Arg His
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<210> 11
 <211> 27
 <212> DNA
 <213> Artificial Sequence

<220>

<223> pET22b(+) forward oligonucleotide primer for
 Tumstatin

<400> 11
 cgggatccgg gtttgaaagg aaaacgt

27

<210> 12
 <211> 27
 <212> DNA
 <213> Artificial Sequence

<220>

<223> pET22b(+) reverse oligonucleotide primer for
 Tumstatin

<400> 12
 cccaagcttt cagtgtcttt tcttcat

27

<210> 13
 <211> 8
 <212> PRT
 <213> Artificial Sequence

<220>

<223> Additional vector sequence added to protein

<400> 13
 Met Asp Ile Gly Ile Asn Ser Asp
 1 5

<210> 14
 <211> 7
 <212> PRT
 <213> Artificial Sequence

<220>

<223> Additional vector sequence added to protein

<400> 14
 Lys Leu Ala Ala Ala Leu Glu
 1 5

<210> 15
 <211> 28
 <212> DNA
 <213> Artificial Sequence

<220>

<223> pPICZaA forward oligonucleotide primer for
 Arresten

<400> 15
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<210> 16

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<211> 35
<212> DNA
<213> Artificial Sequence

<220>
<223> pPICZaA reverse oligonucleotide primer for
Arresten

<400> 16
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<210> 17
<211> 31
<212> DNA
<213> Artificial Sequence

<220>
<223> pPICZaA forward oligonucleotide primer for
Canstatin

<400> 17
ttcgggaattc gtcagcatcg gctacctcct g

31

<210> 18
<211> 32
<212> DNA
<213> Artificial Sequence

<220>
<223> pPICZaA reverse oligonucleotide primer for
Canstatin

<400> 18
gggggtacccc caggttcttc atgcacacct gg

32

<210> 19
<211> 244
<212> PRT
<213> Artificial Sequence

<220>
<223> Tumstatin (amino acids 1-244)

<400> 19
Pro Gly Leu Lys Gly Lys Arg Gly Asp Ser Gly Ser Pro Ala Thr Trp
1 5 10 15
Thr Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile
20 25 30
Pro Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe
35 40 45
Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr
50 55 60
Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys
65 70 75 80
Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr
85 90 95
Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr
100 105 110
Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly

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	115					120					125				
Pro	Ala	Ile	Ala	Ile	Ala	Val	His	Ser	Gln	Thr	Thr	Asp	Ile	Pro	Pro
	130					135					140				
Cys	Pro	His	Gly	Trp	Ile	Ser	Leu	Trp	Lys	Gly	Phe	Ser	Phe	Ile	Met
145					150					155					160
Phe	Thr	Ser	Ala	Gly	Ser	Glu	Gly	Thr	Gly	Gln	Ala	Leu	Ala	Ser	Pro
				165					170					175	
Gly	Ser	Cys	Leu	Glu	Glu	Phe	Arg	Ala	Ser	Pro	Phe	Leu	Glu	Cys	His
			180					185				190			
Gly	Arg	Gly	Thr	Cys	Asn	Tyr	Tyr	Ser	Asn	Ser	Tyr	Ser	Phe	Trp	Leu
	195					200					205				
Ala	Ser	Leu	Asn	Pro	Glu	Arg	Met	Phe	Arg	Lys	Pro	Ile	Pro	Ser	Thr
	210				215						220				
Val	Lys	Ala	Gly	Glu	Leu	Glu	Lys	Ile	Ile	Ser	Arg	Cys	Gln	Val	Cys
225					230					235					240
Met	Lys	Lys	Arg												

<210> 20
<211> 124
<212> PRT
<213> Artificial Sequence

<220>
<223> Tumstatin 333 (amino acids 2-125 of SEQ ID NO:10)

<400> 20
Gly Leu Lys Gly Lys Arg Gly Asp Ser Gly Ser Pro Ala Thr Trp Thr
1 5 10 15
Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile Pro
20 25 30
Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe Leu
35 40 45
Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr Leu
50 55 60
Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys Asn
65 70 75 80
Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr Trp
85 90 95
Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr Gly
100 105 110
Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val
115 120

<210> 21
<211> 119
<212> PRT
<213> Artificial Sequence

<220>
<223> Tumstatin 334 (amino acids 126-244 of SEQ ID NO:10)

<400> 21
Cys Glu Gly Pro Ala Ile Ala Ile Ala Val His Ser Gln Thr Thr Asp
1 5 10 15
Ile Pro Pro Cys Pro His Gly Trp Ile Ser Leu Trp Lys Gly Phe Ser

[illegible]

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<220>
<223> Tum-1 (Tumstatin N53) (amino acids 54-244 of SEQ
ID NO:10)
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<210> 23
<211> 132
<212> PRT
<213> Artificial Sequence
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<220>
<223> Tum-2 (amino acids 1-132 of SEQ ID NO:10)

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<400> 23

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Pro Gly Leu Lys Gly Lys Arg Gly Asp Ser Gly Ser Pro Ala Thr Trp
 1          5          10          15
Thr Thr Arg Gly Phe Val Phe Thr Arg His Ser Gln Thr Thr Ala Ile
          20          25          30
Pro Ser Cys Pro Glu Gly Thr Val Pro Leu Tyr Ser Gly Phe Ser Phe
          35          40          45
Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr
          50          55          60
Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys
65          70          75          80
Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr
          85          90          95
Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr
          100          105          110
Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly
          115          120          125
Pro Ala Ile Ala
          130
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<210> 24

<211> 112

<212> PRT

<213> Artificial Sequence

<220>

<223> Tum-3 (amino acids 133-244 of SEQ ID NO:10)

<400> 24

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Ile Ala Val His Ser Gln Thr Thr Asp Ile Pro Pro Cys Pro His Gly
 1          5          10          15
Trp Ile Ser Leu Trp Lys Gly Phe Ser Phe Ile Met Phe Thr Ser Ala
          20          25          30
Gly Ser Glu Gly Thr Gly Gln Ala Leu Ala Ser Pro Gly Ser Cys Leu
          35          40          45
Glu Glu Phe Arg Ala Ser Pro Phe Leu Glu Cys His Gly Arg Gly Thr
          50          55          60
Cys Asn Tyr Tyr Ser Asn Ser Tyr Ser Phe Trp Leu Ala Ser Leu Asn
65          70          75          80
Pro Glu Arg Met Phe Arg Lys Pro Ile Pro Ser Thr Val Lys Ala Gly
          85          90          95
Glu Leu Glu Lys Ile Ile Ser Arg Cys Gln Val Cys Met Lys Lys Arg
          100          105          110
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<210> 25

<211> 64

<212> PRT

<213> Artificial Sequence

<220>

<223> Tum-4 (amino acids 181-244 of SEQ ID NO:10)

<400> 25

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Glu Glu Phe Arg Ala Ser Pro Phe Leu Glu Cys His Gly Arg Gly Thr
 1          5          10          15
Cys Asn Tyr Tyr Ser Asn Ser Tyr Ser Phe Trp Leu Ala Ser Leu Asn
          20          25          30
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Pro Glu Arg Met Phe Arg Lys Pro Ile Pro Ser Thr Val Lys Ala Gly
35 40 45
Glu Leu Glu Lys Ile Ile Ser Arg Cys Gln Val Cys Met Lys Lys Arg
50 55 60

<210> 26
<211> 79
<212> PRT
<213> Artificial Sequence

<220>
<223> Tum-5 (amino acids 54-132 of SEQ ID NO:10)

<400> 26
Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr Leu Gly Ser Cys Leu
1 5 10 15
Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys Asn Val Asn Asp Val
20 25 30
Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr Trp Leu Ser Thr Pro
35 40 45
Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr Gly Arg Ala Leu Glu
50 55 60
Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly Pro Ala Ile Ala
65 70 75

<210> 27
<211> 20
<212> PRT
<213> Artificial Sequence

<220>
<223> T1 (amino acids 1-20 of SEQ ID NO:10)

<400> 27
Pro Gly Leu Lys Gly Lys Arg Gly Asp Ser Gly Ser Pro Ala Thr Trp
1 5 10 15
Thr Thr Arg Gly
20

<210> 28
<211> 20
<212> PRT
<213> Artificial Sequence

<220>
<223> T2 (amino acids 54-73 of SEQ ID NO:10)

<400> 28
Asn Gln Arg Ala His Gly Gln Asp Leu Gly Thr Leu Gly Ser Cys Leu
1 5 10 15
Gln Arg Phe Thr
20

<210> 29
<211> 20

<212> PRT
 <213> Artificial Sequence

<220>
 <223> T3 (amino acids 69-88 of SEQ ID NO:10)

<400> 29
 Leu Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys Asn Val Asn Asp
 1 5 10 15
 Val Cys Asn Phe
 20

<210> 30
 <211> 20
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T4 (amino acids 84-103 of SEQ ID NO:10)

<400> 30
 Asp Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser Tyr Trp Leu Ser
 1 5 10 15
 Thr Pro Ala Leu
 20

<210> 31
 <211> 19
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T5 (amino acids 99-117 of SEQ ID NO:10)

<400> 31
 Ser Thr Pro Ala Leu Met Pro Met Asn Met Ala Pro Ile Thr Gly Arg
 1 5 10 15
 Ala Leu Glu

<210> 32
 <211> 19
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T6 (amino acids 114-132 of SEQ ID NO:10)

<400> 32
 Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr Val Cys Glu Gly Pro
 1 5 10 15
 Ala Ile Ala

<210> 33

<211> 88
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Tumstatin-45-132 (amino acids 45-132 of SEQ ID
 NO:10)

<400> 33
 Gly Phe Ser Phe Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln
 1 5 10 15
 Asp Leu Gly Thr Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro
 20 25 30
 Phe Leu Phe Cys Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn
 35 40 45
 Asp Tyr Ser Tyr Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met
 50 55 60
 Ala Pro Ile Thr Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr
 65 70 75 80
 Val Cys Glu Gly Pro Ala Ile Ala
 85

<210> 34
 <211> 88
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Tumstatin-5-126-C-A (amino acids 45-132 of SEQ ID
 NO:10; alanine has been substituted for the
 cysteine residue at position 126 of the
 full-length Tumstatin molecule)

<400> 34
 Gly Phe Ser Phe Leu Phe Val Gln Gly Asn Gln Arg Ala His Gly Gln
 1 5 10 15
 Asp Leu Gly Thr Leu Gly Ser Cys Leu Gln Arg Phe Thr Thr Met Pro
 20 25 30
 Phe Leu Phe Cys Asn Val Asn Asp Val Cys Asn Phe Ala Ser Arg Asn
 35 40 45
 Asp Tyr Ser Tyr Trp Leu Ser Thr Pro Ala Leu Met Pro Met Asn Met
 50 55 60
 Ala Pro Ile Thr Gly Arg Ala Leu Glu Pro Tyr Ile Ser Arg Cys Thr
 65 70 75 80
 Val Ala Glu Gly Pro Ala Ile Ala
 85

<210> 35
 <211> 9
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> synthetic blocking peptide

<400> 35
 Cys Asp Cys Arg Gly Asp Cys Phe Cys

1

5

<210> 36
 <211> 5
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> synthetic blocking peptide

<400> 36
 Cys Asn Gly Arg Cys
 1 5

<210> 37
 <211> 25
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T7 (amino acids 74-98 of SEQ ID NO:10)

<400> 37
 Thr Met Pro Phe Leu Phe Cys Asn Val Asn Asp Val Cys Asn Phe Ala
 1 5 10 15
 Ser Arg Asn Asp Tyr Ser Tyr Trp Leu
 20 25

<210> 38
 <211> 25
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T7-mutant (amino acids 74-98 of SEQ ID NO:10;
 methionine has been substituted for the leucine
 residue at position 78 of the full-length
 Tumstatin molecule, and isoleucine has been
 substituted for valine at position 82, and
 asparagine has been substituted for aspartic acid
 at position 84)

<400> 38
 Thr Met Pro Phe Met Phe Cys Asn Ile Asn Asn Val Cys Asn Phe Ala
 1 5 10 15
 Ser Arg Asn Asp Tyr Ser Tyr Trp Leu
 20 25

<210> 39
 <211> 27
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> T8 (amino acids 69-95 of SEQ ID NO:10; lysine has

been substituted for the leucine residue at
position 69 of the full-length Tumstatin molecule)

<400> 39

```
Lys Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Cys Asn Val Asn Asp
 1           5           10           15
Val Cys Asn Phe Ala Ser Arg Asn Asp Tyr Ser
          20           25
```

<210> 40

<211> 27

<212> PRT

<213> Artificial Sequence

<220>

<223> T8-3 (amino acids 69-95 of SEQ ID NO:10; lysine
has been substituted for the leucine residue at
position 69 of the full-length Tumstatin molecule,
and serine has been substituted for the cysteine
residues at positions 80 and 86)

<400> 40

```
Lys Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Ser Asn Val Asn Asp
 1           5           10           15
Val Ser Asn Phe Ala Ser Arg Asn Asp Tyr Ser
          20           25
```

<210> 41

<211> 19

<212> PRT

<213> Artificial Sequence

<220>

<223> TP3 (amino acids 77-95 of SEQ ID NO:10; lysine has
been substituted for the phenylalanine residue at
position 77 of the full-length Tumstatin molecule,
and cysteine has been substituted for the aspartic
acid at position 84)

<400> 41

```
Lys Leu Phe Cys Asn Val Asn Cys Val Cys Asn Phe Ala Ser Arg Asn
 1           5           10           15
Asp Tyr Ser
```

<210> 42

<211> 27

<212> PRT

<213> Artificial Sequence

<220>

<223> P2 (amino acids 69-95 of SEQ ID NO:10; lysine has
been substituted for the leucine residue at
position 69 of the full-length Tumstatin molecule,
and aspartic acid has been substituted for the
cysteine residues at positions 80 and 86)

Lys Gln Arg Phe Thr Thr Met Pro Phe Leu Phe Asp Asn Val Asn Asp
1 5 10 15
Val Asp Asn Phe Ala Ser Arg Asn Asp Tyr Ser
20 25

<213> Artificial Sequence

<223> Scrambled peptide SP1

Ala Asn Met Ser Arg Asn Val Phe Phe Asp Cys Thr Ser Phe Pro Val
1 5 10 15
Cys Gln Lys Phe Leu Asn Asp Thr Arg Asn Tyr
20 25

<213> Artificial Sequence

<223> Scrambled peptide SP2

Thr Phe Asn Cys Val Lys Asn Tyr Gln Arg Leu Asp Phe Thr Ser Arg
1 5 10 15
Phe Val Met Asp Ser Cys Ala Asn Phe Pro Asn
20 25

<213> rtificial Sequence

<223> Generic peptide

<223> X at position 14 is a hydrogen or a peptidyl chain
of 1 to 12 amino acids

<400> 45

Xaa Xaa Leu Phe Xaa Asn Val Asn Xaa Val Xaa Asn Phe Xaa
 1 5 10

<210> 46

<211> 4

<212> PRT

<213> Artificial Sequence

<220>

<223> Generic peptide

<400> 46

Thr Thr Met Pro
 1

<210> 47

<211> 5

<212> PRT

<213> Artificial Sequence

<220>

<223> Generic peptide

<400> 47

Phe Thr Thr Met Pro
 1 5

<210> 48

<211> 6

<212> PRT

<213> Artificial Sequence

<220>

<223> Generic peptide

<400> 48

Arg Phe Thr Thr Met Pro
 1 5

<210> 49

<211> 7

<212> PRT

<213> Artificial Sequence

<220>

<223> Generic peptide

<400> 49

Gln Arg Phe Thr Thr Met Pro
 1 5

<210> 50

<211> 8

<212> PRT
 <213> Artificial Sequence

<220>
 <223> Generic peptide

<400> 50
 Leu Gln Arg Phe Thr Thr Met Pro
 1 5

<210> 51
 <211> 8
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Generic peptide

<400> 51
 Lys Gln Arg Phe Thr Thr Met Pro
 1 5

<210> 52
 <211> 4
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Generic peptide

<400> 52
 Ala Ser Arg Asn
 1

<210> 53
 <211> 5
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Generic peptide

<400> 53
 Ala Ser Arg Asn Asp
 1 5

<210> 54
 <211> 6
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> Generic peptide

<400> 54

Ala Ser Arg Asn Asp Tyr
1 5

<210> 55
<211> 7
<212> PRT
<213> Artificial Sequence

<220>
<223> Generic peptide

<400> 55
Ala Ser Arg Asn Asp Tyr Ser
1 5

<210> 56
<211> 8
<212> PRT
<213> Artificial Sequence

<220>
<223> Generic peptide

<400> 56
Ala Ser Arg Asn Asp Tyr Ser Tyr
1 5

<210> 57
<211> 9
<212> PRT
<213> Artificial Sequence

<220>
<223> Generic peptide

<400> 57
Ala Ser Arg Asn Asp Tyr Asp Tyr Trp
1 5

<210> 58
<211> 10
<212> PRT
<213> Artificial Sequence

<220>
<223> Generic peptide

<400> 58
Ala Ser Arg Asn Asp Tyr Ser Tyr Trp Leu
1 5 10